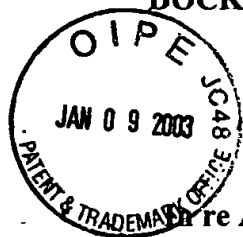


DOCKET NO.: ISIS-3455

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE



In re Application of:

Cowsert *et al.*

Confirmation No.: 7206

Serial No.: 09/295,463

Group Art Unit: 1631

Filing Date: April 13, 1999

Examiner: Marschel

For: IDENTIFICATION OF GENETIC TARGETS FOR MODULATION BY  
OLIGONUCLEOTIDES AND GENERATION OF OLIGONUCLEOTIDES FOR  
GENE MODULATION

EXPRESS MAIL LABEL NO: EL884781742US  
DATE OF DEPOSIT: January 9, 2003

Assistant Commissioner for Patents  
Washington DC 20231

Sir:

**REPLY PURSUANT TO 37 CFR § 1.114**

This is in response to the Advisory Action mailed November 29, 2002 and to the Final Office Action mailed September 10, 2002, and is being filed along with a request for continued examination (RCE) in connection with the above-identified patent application. Applicants respectfully request that the application be amended as follows.

**In the Claims:**

Please amend claim 55 to read as follows:

55. (Amended three times) A method comprising:

generating *in silico* virtual compounds according to thermodynamic property, target accessibility, targeting to functional regions of target nucleic acid sequence, or uniform distribution to target nucleic acid sequence, wherein synthetic compounds corresponding to said virtual compounds modulate the expression of a target nucleic acid sequence;

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EL884781742US

Box ☒ AF  
☐ ISSUE FEE

Assistant Commissioner for Patents  
Washington DC 20231

Sir:

### REQUEST FOR CONTINUED EXAMINATION (RCE) TRANSMITTAL

This is a Request for Continued Examination (RCE) under 37 CFR § 1.114 of the above-identified application.

1. Submission required under 37 CFR § 1.114

- ☐ Previously submitted
- filed on ☐ Consider the amendment(s)/reply under 37 CFR § 1.116 previously  
(Any unentered amendment(s) referred to above will be entered).
- filed on ☐ Consider the arguments in the Appeal Brief or Reply Brief previously
- ☐ Other:
- ☒ Enclosed

01/14/2003 MGBREM1 00000005 09295463

01 FC:2801

375.00 OP

01/14/2003 MGBREM1 00000005 09295463

55.00 OP

02 FC:2251

- ☒ Amendment/Reply
- ☐ Affidavit(s)/Declaration(s)
- ☐ Information Disclosure Statement (IDS)
- ☐ Other:

## 2. Miscellaneous

☐ Suspension of action on the above-identified application is requested under 37 CFR §1.103(c) for a period of \_\_\_\_\_ months. (Period of suspension shall not exceed three (3) months; Fee under 37 CFR §1.17(i) required)

**FEE CALCULATION:**

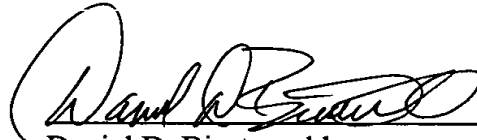
				SMALL ENTITY		NOT SMALL ENTITY	
<input checked="" type="checkbox"/> RCE BASIC FILING FEE				\$375.00	\$375.00	\$750.00	\$
	CLAIMS REMAINING AFTER RCE	HIGHEST NO. PAID FOR	EXTRA				
TOTAL CLAIMS	35	54 (20 MINIMUM)	0	\$9 EACH	\$0.00	\$18 EACH	\$
INDEP. CLAIMS	29	45 (3 MINIMUM)	0	\$42 EACH	\$0.00	\$84 EACH	\$
FIRST PRESENTATION OF MULTIPLE DEPENDENT				\$140	\$0.00	\$290	\$
<input checked="" type="checkbox"/> ONE MONTH EXTENSION OF TIME				\$55	\$55.00	\$110	\$
<input type="checkbox"/> TWO MONTH EXTENSION OF TIME				\$205	\$0.00	\$410	\$
<input type="checkbox"/> THREE MONTH EXTENSION OF TIME				\$465	\$0.00	\$930	\$
<input type="checkbox"/> FOUR MONTH EXTENSION OF TIME				\$725	\$0.00	\$1450	\$
<input type="checkbox"/> FIVE MONTH EXTENSION OF TIME (May only be taken when filing RCE in lieu of Appeal Brief)				\$985	\$0.00	\$1970	\$
<input type="checkbox"/> LESS ANY EXTENSION FEE ALREADY PAID				minus	(\$0.00)	minus	(\$ )
<input type="checkbox"/> TERMINAL DISCLAIMER				\$55	\$0.00	\$110	\$
TOTAL FEE DUE					\$430.00		\$

- ☒ Two checks equaling the amount of **\$430.00** are attached. Please charge any deficiency or credit any overpayment to Deposit Account No. 23-3050.
- ☐ Please charge Deposit Account No. 23-3050 in the amount of \$ \_\_\_\_\_.00. This sheet is attached in duplicate.
- ☒ A response to the **Final Office Action** dated **September 10, 2002** was due on **December 10, 2002**. Petition is hereby made under 37 CFR § 1.136(a) to extend the time period for response from **December 10, 2002** to and through **January 10, 2003** comprising an extension of the shortened statutory period of **ne** month.

- ☒ The Commissioner is hereby requested to grant an extension of time for the appropriate length of time, should one be necessary, in connection with this filing or any future filing submitted to the U.S. Patent and Trademark Office in the above-identified application during the pendency of this application. The Commissioner is further authorized to charge any fees related to any such extension of time to Deposit Account No. 23-3050. This sheet is provided in duplicate.
- ☒ The Commission is hereby authorized to charge payment of the above fees associated with this communication or credit any overpayment to Deposit Account No. 23-3050. This sheet is attached in duplicate.

**SHOULD ANY DEFICIENCIES APPEAR** with respect to this application, including deficiencies in payment of fees, missing parts of the application or otherwise, the United States Patent and Trademark Office is respectfully requested to promptly notify the undersigned.

Date: January 9, 2003



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